SEP 0 8 2003

SEQUENCE LISTING

ARCANGEL, Phillip CHIEN, David Y.

<120> HCV ASSAY

<130> 2300~19199

<150> 60/409,515

<151> 2002-09-09

<160> 9

<170> PatentIn version 3.2

<210> 1

<211> 2058

<212> DNA

<213> Artificial

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<223> NS3/41 conformational epitope DNA sequence

<400> 1

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Met Ala Pro Ile Thr Ala Tyr Ala Gln Gln Thr Arg Gly Leu Leu Gly
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gag gtc cag att gtg tca act gct gcc caa acc ttc ctg gca acg tgc 144 Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr Phe Leu Ala Thr Cys

atc aat ggg gtg tgc tgg act gtc tac cac ggg gcc gga acg agg acc 192
Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg Thr
50 55 60

atc gcg tca ccc aag ggt cct gtc atc cag atg tat acc aat gta gac 240

Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met Tyr Thr Asn Val Asp
65 70 75 80

caa gac ctt gtg ggc tgg ccc gct ccg caa ggt agc cga tca ttg aca 288 Gln Asp Leu Val Gly Trp Pro Ala Pro Gln Gly Ser Arg Ser Leu Thr

ccc tgc act tgc ggc tcc tcg gac ctt tac ctg gtc acg agg cac gcc 336
Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg His Ala
100 105 110

gat gtc att ccc gtg cgc cgg cgg ggt gat agc agg ggc agc ctg ctg 384
Asp Val Ile Pro Val Arg Arg Arg Gly Asp Ser Arg Gly Ser Leu Leu
115

tcg ccc cgg ccc att tcc tac ttg aaa ggc tcc tcg ggg ggt ccg ctg 432 Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro Leu 130 135 140

| _ | ~ | | _ | | | _ | | | | | | _ | | gtg Val | _ | 480 |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------------------|---|------|
| | | | | | | | | | | | | | | aac Asn 175 | | 528 |
| | | | _ | | | | | | _ | _ | | | | cca Pro | | 576 |
| _ | | | | _ | | _ | | _ | | | | _ | | aca Thr | | 624 |
| _ | | | _ | | _ | _ | _ | _ | _ | | - | _ | _ | Gly | | 672 |
| - | | | _ | | | | | _ | _ | _ | | _ | | ttt Phe | | 720 |
| ~ | | _ | | _ | - | | | | _ | | | | | acc Thr 255 | | 768 |
| | | | | | | | | | | | | | | tac Tyr | | 816 |
| _ | | | - | - | | | | | | | | | | ata Ile | | 864 |
| | _ | _ | | _ | | | _ | - | - | | | | _ | ggc Gly | | 912 |
| | | | | | | | | | | | | | | gtt Val | | 960 |
| | | | | | | | | | | | | | | ccc Pro 335 | | 1008 |
| | | | _ | - | _ | | | | | | | | | tac Tyr | | 1056 |
| | | | | | | | | | | | | | | atc Ile | | 1104 |
| | | | | | | | | | | | | | | gtc Val | | 1152 |

| | tc aat gcc le Asn Ala | | | | Leu Asp | | |) |
|-----------|---------------------------------|---|---------|---------|---------|-----------|---|---|
| | cc atc ggc ro Ile Gly 405 | | | | | | | 3 |
| | at acc ggc yr Thr Gly 420 | | | Val Ile | | | | 5 |
| Val Thr G | ag aca gtc ln Thr Val 35 | - | _ | _ | | Thr Ile | | 4 |
| | cg ctc ccc hr Leu Pro | | | | | | | 2 |
| | gc agg ggg ly Arg Gly | - | | _ | Phe Val | | | 0 |
| ~ ~ | cc tcc ggc ro Ser Gly 485 | - | | - | _ | | | 8 |
| | gc tgt gct ly Cys Ala 500 | | | Thr Pro | | | | 6 |
| Arg Leu A | ga gcg tac rg Ala Tyr 15 | - | _ | | | Cys Gln | - | 4 |
| | aa ttt tgg lu Phe Trp | | | | | | | 2 |
| | tt cta tcc he Leu Ser | | | | Glu Asn | | | 0 |
| | cg tac caa la Tyr Gln 565 | | | | | | | 8 |
| | gg gac cag rp Asp Gln 580 | | | Leu Ile | - | | | 6 |
| Leu His G | gg cca aca ly Pro Thr 95 | _ | | | | . Val Gln | | 4 |
| _ | cc ctg acg hr Leu Thr | | Val Thr | | _ | _ | - | 2 |

. .

| | | g agc acc tgg gtg r Ser Thr Trp Val 635 | | 920 |
|---|---|---|-----|-----|
| Val Leu Ala Ala I | | tgc ctg tca aca r Cys Leu Ser Thr 650 | | 968 |
| | | c ggg aag ccg gca r Gly Lys Pro Ala 665 | _ | 016 |
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Cys Ile Ile Thr Ser Leu Thr Gly Arg Asp Lys Asn Gln Val Glu Gly 20 25 30

Glu Val Gln Ile Val Ser Thr Ala Ala Gln Thr Phe Leu Ala Thr Cys 35 40 45

Ile Asn Gly Val Cys Trp Thr Val Tyr His Gly Ala Gly Thr Arg Thr 50 60

Ile Ala Ser Pro Lys Gly Pro Val Ile Gln Met Tyr Thr Asn Val Asp 65 70 75 80

Gln Asp Leu Val Gly Trp Pro Ala Pro Gln Gly Ser Arg Ser Leu Thr 85 90 95

Pro Cys Thr Cys Gly Ser Ser Asp Leu Tyr Leu Val Thr Arg Hiś Ala 100 . 105 110

Asp Val Ile Pro Val Arg Arg Gly Asp Ser Arg Gly Ser Leu Leu 115 120 125

Ser Pro Arg Pro Ile Ser Tyr Leu Lys Gly Ser Ser Gly Gly Pro Leu 130 135 140

Leu Cys Pro Ala Gly His Ala Val Gly Ile Phe Arg Ala Ala Val Cys 145 150 155 160

Thr Arg Gly Val Ala Lys Ala Val Asp Phe Ile Pro Val Glu Asn Leu 165 170 175

- Glu Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro Pro 180 185 190
- Val Val Pro Gln Ser Phe Gln Val Ala His Leu His Ala Pro Thr Gly
 195 200 205
- Ser Gly Lys Ser Thr Lys Val Pro Ala Ala Tyr Ala Ala Gln Gly Tyr
- Lys Val Leu Val Leu Asn Pro Ser Val Ala Ala Thr Leu Gly Phe Gly 225 230 235 240
- Ala Tyr Met Ser Lys Ala His Gly Ile Asp Pro Asn Ile Arg Thr Gly 245 250 255
- Val Arg Thr Ile Thr Thr Gly Ser Pro Ile Thr Tyr Ser Thr Tyr Gly 260 265 270
- Lys Phe Leu Ala Asp Gly Gly Cys Ser Gly Gly Ala Tyr Asp Ile Ile 275 280 285
- Ile Cys Asp Glu Cys His Ser Thr Asp Ala Thr Ser Ile Leu Gly Ile 290 295 300
- Gly Thr Val Leu Asp Gln Ala Glu Thr Ala Gly Ala Arg Leu Val Val 305 310 315
- Leu Ala Thr Ala Thr Pro Pro Gly Ser Val Thr Val Pro His Pro Asn 325 330 335
- Ile Glu Glu Val Ala Leu Ser Thr Thr Gly Glu Ile Pro Phe Tyr Gly 340 345 350
- Lys Ala Ile Pro Leu Glu Val Ile Lys Gly Gly Arg His Leu Ile Phe 355 360 365
- Cys His Ser Lys Lys Lys Cys Asp Glu Leu Ala Ala Lys Leu Val Ala 370 380
- Leu Gly Ile Asn Ala Val Ala Tyr Tyr Arg Gly Leu Asp Val Ser Val 385 390 395
- Ile Pro Pro Ile Gly Asp Val Val Val Ala Thr Asp Ala Leu Met 405 410 410
- Thr Gly Tyr Thr Gly Asp Phe Asp Ser Val Ile Asp Cys Asn Thr Cys 420 425 430
- Val Thr Gln Thr Val Asp Phe Ser Leu Asp Pro Thr Phe Thr Ile Glu 435 440 445
- Thr Ile Thr Leu Pro Gln Asp Ala Val Ser Arg Thr Gln Arg Arg Gly
 450 455 460
- Arg Thr Gly Arg Gly Lys Pro Gly Ile Tyr Arg Phe Val Ala Pro Gly 465 470 475 480
- Glu Arg Pro Ser Gly Met Phe Asp Ser Ser Val Leu Cys Glu Cys Tyr 485 490 495

| Asp | Ala | Gly | Cys 500 | Ala | Trp | Tyr | Glu | Leu 505 | Thr | Pro | Ala | Glu | Thr 510 | Thr | Val | |
|------------------------------|------------------|---------------------------|------------------|------------|------------|------------|------------|------------------|------------|------------|------------|------------|------------------|------------|------------------|-----|
| Arg | Leu | Arg 515 | Ala | Tyr | Met | Asn | Thr 520 | Pro | Gly | Leu | Pro | Val 525 | Cys | Gln | Asp | |
| His | Leu 530 | Glu | Phe | Trp | | Gly 535 | Val | Phe | Thr | Gly | Leu 540 | Thr | His | Ile | Asp | |
| Ala 545 | His | Phe | Leu | Ser | Gln 550 | Thr | Lys | Gln | Ser | Gly 555 | Glu | Asn | Leu | Pro | Tyr 560 | |
| Leu | Val | Ala | Tyr | Gln 565 | Ala | Thr | Val | Сув | Ala 570 | Arg | Ala | Gln | Ala | Pro 575 | Pro | |
| Pro | Ser | Trp | Asp 580 | Gln | Met | Trp | Lys | Cys 585 | Leu | Ile | Arg | Leu | Lys 590 | Pro | Thr | |
| Leu | His | Gly 595 | Pro | Thr | Pro | Leu | Leu 600 | Tyr | Arg | Leu | Gly | Ala 605 | Val | Gln | Asn _. | |
| Glu | Ile 610 | Thr | Leu | Thr | His | Pro 615 | Val | Thr | Lys | Tyr | Ile 620 | Met | Thr | Сув | Met | |
| Ser 625 | Ala | Asp | Leu | Glu | Val 630 | Val | Thr | Ser | Thr | Trp 635 | Val | Leu | Val | Gly | Gly 640 | |
| Val | Leu | Ala | Ala | Leu 645 | Ala | Ala | Tyr | Cys | Leu 650 | Ser | Thr | Gly | Суѕ | Val 655 | Val | |
| Ile | Val | Gly | Arg 660 | Val | Val | Leu | Ser | Gly 665 | Lys | Pro | Ala | Ile | Ile 670 | Pro | Asp | |
| Arg | Glu | Val 675 | Leu | Tyr | Arg | Glu | Phe 680 | Asp | Glu | Met | Glu | Glu 685 | Cys | | | |
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| <220 <223 | | MEFA | 12 I | DNA : | seque | ence | | | | | | | | | | |
| <400 | | 3 | 225 | aat | ~++ | +~+ | a++ | ++~ | 224 | ~~+ | ~~~ | ~~~ | | ~ | ~ | 40 |
| _ | gct Ala | | - | - | _ | - | - | _ | _ | | _ | | | _ | | 48 |
| ggt Gly | att Ile | att Ile | aac Asn 20 | ttc Phe | gag Glu | cag Gln | aag Lys | gaa Glu 25 | agt Ser | aat Asn | gga Gly | cca Pro | gtg Val 30 | aag Lys | gtg Val | 96 |
| | gga Gly | | | | | | | | | | | | | | | 144 |
| | gag Glu 50 | | | | | | | | | | | | | | | 192 |

| | aat Asn | | | | _ | _ | | _ | | _ | | | | | | 240 |
|------------|-------------------|------------|------------|------------|------------|-------------------|------------|------------|------------|------------|-------------------|------------|------------|------------|------------|-----|
| | ata Ile | _ | | | _ | _ | _ | | _ | | | | - | _ | _ | 288 |
| | acc Thr | | | | | | | | | | | | | | | 336 |
| _ | act Thr | | _ | _ | | | - | _ | _ | _ | | _ | | | ~ | 384 |
| | tct Ser 130 | _ | | | | | _ | _ | | | | | _ | | | 432 |
| _ | acg Thr | - | | | | | | - | | | _ | _ | | _ | | 480 |
| | cac His | | | | | | | | | | | | | | | 528 |
| _ | gca Ala | | | - | _ | | | _ | | | - | | | | | 576 |
| | gct Ala | _ | | | | | | - | | _ | | | _ | | | 624 |
| | gat Asp 210 | | | | | | | | | | | | | | | 672 |
| | atc Ile | _ | | | | | | _ | | | _ | _ | | | ~ | 720 |
| | GJA aaa | | | | | | | | | | | | | | | 768 |
| | gcc Ala | | | | | | | | | | | | | | | 816 |
| | gcg Ala | | _ | | _ | _ | | | - | | _ | | | _ | | 864 |
| tcc Ser | gtc Val 290 | act Thr | gtg Val | ccc Pro | cat His | ccc Pro 295 | aac Asn | atc Ile | gag Glu | gag Glu | gtt Val 300 | gct Ala | ctg Leu | tcc Ser | acc Thr | 912 |

| | | | | | | | | | | | | ctc Leu | | | | 960 | |
|------------|------------|------------|-------------------|------------|------------|------------|------------|-------------------|------------|------------|------------|-------------------|-------------------|------------|------------|------|--|
| | | | | | | | | | | | | aag Lys | | | | 1008 | |
| gaa Glu | ctc Leu | gcc Ala | gca Ala 340 | aag Lys | ctg Leu | gtc Val | gca Ala | ttg Leu 345 | ggc Gly | atc Ile | aat Asn | gcc Ala | gtg Val 350 | gcc Ala | tac Tyr | 1056 | |
| | _ | | | - | | | - | | | | _ | ggc Gly 365 | | | - | 1104 | |
| | | | | | | | | | | | | ggc Gly | | | | 1152 | |
| | | | | | | | | | | | | aag Lys | | | | 1200 | |
| | | | | | | | | | | | | gag Glu | | | | 1248 | |
| | | | | | | | | | | | | atg Met | | | | 1296 | |
| | | | | | | | | | | | | ggc Gly 445 | | | | 1344 | |
| | | | | | | | | | | | | gat Asp | | | | 1392 | |
| | _ | | | _ | _ | | | | _ | | _ | cag Gln | _ | | - | 1440 | |
| | - | | _ | | | _ | | | _ | | - | aat Asn | _ | | gtg Val | 1488 | |
| | | | | | | | | | | | | ttt Phe | | | atg Met | 1536 | |
| _ | _ | _ | _ | | | _ | _ | | | | _ | 999 Gly 525 | _ | | atg Met | 1584 | |
| | | | | | | | | | | | | | | | cgc Arg | 1632 | |

| | cgg Arg 545 | | | | cct Pro | | | | | | | | | | | | 1680 |
|---|-------------------|---|---|---|-------------------|---|---|---|---|---|---|---|---|---|---|---|------|
| | | | | | tcc Ser 565 | | | | | | | | | | | | 1728 |
| | | | | | cgg Arg | | | | | | | | | | | | 1776 |
| | ~ | - | | | ccc Pro | _ | | | | _ | | | _ | | _ | | 1824 |
| | | | | | gtc Val | | | | | | | | | | | | 1872 |
| | | | _ | | gcg Ala | | _ | _ | | | | _ | | | | - | 1920 |
| | | | _ | | gac Asp 645 | | _ | | | _ | _ | | | _ | _ | | 1968 |
| | | - | | | aac Asn | | | _ | _ | _ | _ | _ | | _ | | | 2016 |
| | | _ | | _ | ggt Gly | | _ | | _ | _ | _ | | | | | - | 2064 |
| | _ | | | | gcg Ala | - | _ | _ | | | | | | - | - | _ | 2112 |
| | | | | | agg Arg | | | | | | _ | | | | | | 2160 |
| | | | | | gac Asp 725 | | | | | | | | | | | | 2208 |
| | | | | | cca Pro | - | _ | | | _ | | | | | | _ | 2256 |
| | | | | | ttc Phe | | | | | | | | | | | | 2304 |
| · | _ | _ | _ | _ | agg Arg | | | _ | _ | | | | | _ | _ | _ | 2352 |
| | | | | | | | | | | , | | | | | | | |

act tcc cct atc ccc aag gct cgt cgg ccc gag ggc agg acc tgg gct 2400 Thr Ser Pro Ile Pro Lys Ala Arg Arg Pro Glu Gly Arg Thr Trp Ala 785 790 795 800

cag ccc ggt tac cct tgg ccc ctc tat ggc aat aag gac aga cgg tct 2448 Gln Pro Gly Tyr Pro Trp Pro Leu Tyr Gly Asn Lys Asp Arg Arg Ser 805 810 815

aca ggt aag too tgg ggt aag coa ggg tac cot tgg coo taatgagtog ac 2499 Thr Gly Lys Ser Trp Gly Lys Pro Gly Tyr Pro Trp Pro 820 825

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<212> PRT

<213> Artificial

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Gly Ile Ile Asn Phe Glu Gln Lys Glu Ser Asn Gly Pro Val Lys Val 20 25 30

Trp Gly Ser Ile Lys Gly Leu Thr Glu Gly Leu His Gly Phe His Val

His Glu Phe Gly Asp Asn Thr Ala Gly Cys Thr Ser Ala Gly Pro His 50 60

Phe Asn Pro Leu Ser Thr Arg Gly Cys Asn Cys Ser Ile Tyr Pro Gly 65 70 75 80

His Ile Thr Gly His Arg Met Ala Trp Lys Leu Gly Ser Ala Ala Arg

Thr Thr Ser Gly Phe Val Ser Leu Phe Ala Pro Gly Ala Lys Gln Asn 100 105 110

Glu Thr His Val Thr Gly Gly Ala Ala Ala Arg Thr Thr Ser Gly Leu 115 120 125

Thr Ser Leu Phe Ser Pro Gly Ala Ser Gln Asn Ile Gln Leu Ile Thr 130 135 140

Ser Thr Asp Asn Ser Ser Pro Pro Val Val Pro Gln Ser Phe Gln Val 145 150 155 160

Ala His Leu His Ala Pro Thr Gly Ser Gly Lys Ser Thr Lys Val Pro

Ala Ala Tyr Ala Ala Gln Gly Tyr Lys Val Leu Val Leu Asn Pro Ser 180 185 190

Val Ala Ala Thr Leu Gly Phe Gly Ala Tyr Met Ser Lys Ala His Gly
195 200 205

- Ile Asp Pro Asn Ile Arg Thr Gly Val Arg Thr Ile Thr Thr Gly Ser 210 220
- Pro Ile Thr Tyr Ser Thr Tyr Gly Lys Phe Leu Ala Asp Gly Gly Cys
 225 230 235 240
- Ser Gly Gly Ala Tyr Asp Ile Ile Ile Cys Asp Glu Cys His Ser Thr \$245\$
- Asp Ala Thr Ser Ile Leu Gly Ile Gly Thr Val Leu Asp Gln Ala Glu 260 265 270
- Thr Ala Gly Ala Arg Leu Val Val Leu Ala Thr Ala Thr Pro Pro Gly 275 280 285
- Ser Val Thr Val Pro His Pro Asn Ile Glu Glu Val Ala Leu Ser Thr 290 295 300
- Thr Gly Glu Ile Pro Phe Tyr Gly Lys Ala Ile Pro Leu Glu Val Ile 305 310 315 320
- Lys Gly Gly Arg His Leu Ile Phe Cys His Ser Lys Lys Cys Asp 325 330 335
- Glu Leu Ala Ala Lys Leu Val Ala Leu Gly Ile Asn Ala Val Ala Tyr 340 345 350
- Tyr Arg Gly Leu Asp Val Ser Val Ile Pro Thr Ser Gly Asp Val Val 355 360 365
- Val Val Ala Thr Asp Ala Leu Met Thr Gly Tyr Thr Gly Asp Phe Asp 370 375 380
- Ser Val Ile Asp Cys Asn Thr Cys Ala Cys Ser Gly Lys Pro Ala Ile 385 390 395 400
- Ile Pro Asp Arg Glu Val Leu Tyr Arg Glu Phe Asp Glu Met Glu Glu 405 415
- Cys Ser Gln His Leu Pro Tyr Ile Glu Gln Gly Met Met Leu Ala Glu 420 425 430
- Gln Phe Lys Gln Lys Ala Leu Gly Leu Ser Arg Gly Gly Lys Pro Ala 435 440 445
- Ile Val Pro Asp Lys Glu Val Leu Tyr Gln Gln Tyr Asp Glu Met Glu 450 460
- Glu Cys Ser Gln Ala Ala Pro Tyr Ile Glu Gln Ala Gln Val Ile Ala 465 470 475 480
- His Gln Phe Lys Glu Lys Val Leu Gly Leu Ile Asp Asn Asp Gln Val 485 490 495
- Val Val Thr Pro Asp Lys Glu Ile Leu Tyr Glu Ala Phe Asp Glu Met 500 505 510
- Glu Glu Cys Ala Ser Lys Ala Ala Leu Ile Glu Glu Gly Gln Arg Met 515 520 525

Ala Glu Met Leu Lys Ser Lys Ile Gln Gly Leu Leu Gly Ile Leu Arg 530 535 540

Arg His Val Gly Pro Gly Glu Gly Ala Val Gln Trp Met Asn Arg Leu 545 550 560

Ile Ala Phe Ala Ser Arg Gly Asn His Val Ser Pro Thr His Tyr Val
565 570 575

Pro Ser Arg Ser Arg Arg Phe Ala Gln Ala Leu Pro Val Trp Ala Arg 580 585 590

Pro Asp Tyr Asn Pro Pro Leu Val Glu Thr Trp Lys Lys Pro Asp Tyr 595 600 605

Glu Pro Pro Val Val His Gly Arg Ser Ser Arg Arg Phe Ala Gln Ala 610 615 620

Leu Pro Val Trp Ala Arg Pro Asp Tyr Asn Pro Pro Leu Val Glu Thr 625 630 635 640

Trp Lys Lys Pro Asp Tyr Glu Pro Pro Val Val His Gly Arg Lys Thr $645 \hspace{1cm} 650 \hspace{1cm} 655$

Lys Arg Asn Thr Asn Arg Arg Pro Gln Asp Val Lys Phe Pro Gly Gly 660 665 670

Gly Gln Ile Val Gly Gly Val Tyr Leu Leu Pro Arg Arg Gly Pro Arg 675 680 685

Leu Gly Val Leu Ala Thr Arg Lys Thr Ser Pro Ile Pro Lys Ala Arg 690 695 700

Arg Pro Glu Gly Arg Thr Trp Ala Gln Pro Gly Tyr Pro Trp Pro Leu 705 710 715 720

Tyr Gly Asn Lys Asp Arg Arg Ser Thr Gly Lys Ser Trp Gly Lys Pro

Gly Tyr Pro Trp Pro Arg Lys Thr Lys Arg Asn Thr Asn Arg Arg Pro 740 745 750

Gln Asp Val Lys Phe Pro Gly Gly Gln Ile Val Gly Gly Val Tyr 755 760 765

Leu Leu Pro Arg Arg Gly Pro Arg Leu Gly Val Leu Ala Thr Arg Lys
770 775 780

Thr Ser Pro Ile Pro Lys Ala Arg Arg Pro Glu Gly Arg Thr Trp Ala 785 790 795 800

Gln Pro Gly Tyr Pro Trp Pro Leu Tyr Gly Asn Lys Asp Arg Arg Ser 805 810 815

Thr Gly Lys Ser Trp Gly Lys Pro Gly Tyr Pro Trp Pro ; 820 825

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205

200

| | | | | | | | | | | | | | | gcc Ala | | 672 |
|---|---|---|---|---|---|---|---|---|-----|---|---|---|---|-------------------|------------|------|
| | | | | _ | | _ | _ | | | | | | | cta Leu | | 720 |
| | | | | | | | | | | | | | | cca Pro 255 | | 768 |
| | | | | | | | | | | | | | | ggc Gly | | 816 |
| | | | | | | | | | | | | | | tat Tyr | | 864 |
| | | _ | | | | | | | | | | | | ggt Gly | | 912 |
| | | | | | | | | | | | | | | ggg Glý | | 960 |
| | | | | | | | | | | | | | | ggc Gly 335 | | 1008 |
| | | _ | _ | | | | - | | | _ | | - | | ata Ile | | 1056 |
| • | _ | | _ | | | _ | - | _ | | | | _ | | att Ile | | 1104 |
| | _ | | _ | | - | | | | - 3 | | | - | _ | gtg Val | | 1152 |
| | | | | | | | | | | | | | | aac Asn | | 1200 |
| | | | | | | | | | | | | | | ggc Gly 415 | | 1248 |
| _ | | | | _ | _ | | _ | | | _ | | | | ttc Phe | - | 1296 |
| | | - | _ | _ | _ | | _ | | - | _ | _ | _ | _ | - | ttg Leu | 1344 |

| | | | | | | | | | | | gac Asp 460 | | | | | 1392 |
|---|---|---|---|---|---|---|---|---|---|---|-------------------|---|---|---|---|------|
| _ | | _ | | _ | _ | - | _ | | _ | | gat Asp | _ | | _ | | 1440 |
| | | | | _ | | _ | _ | | | _ | tgc Cys | | _ | - | _ | 1488 |
| | _ | | _ | _ | | - | | _ | | | ttc Phe | | | | | 1536 |
| | _ | | | | _ | _ | _ | | _ | | caa Gln | - | | | | 1584 |
| | | | | _ | | | | | _ | | gtg Val 540 | _ | _ | | | 1632 |
| _ | | | | - | | _ | _ | | _ | | tgt Cys | | - | | _ | 1680 |
| | | | | | | | | | | | gag Glu | | | | | 1728 |
| | - | | | _ | | | _ | | | | gtg Val | _ | - | _ | | 1776 |
| | _ | | | | | _ | | | | | act Thr | | | | | 1824 |
| | | | | _ | | - | _ | - | | | aac Asn 620 | | | | | 1872 |
| _ | | | | _ | | | - | _ | | _ | caa Gln | - | | | | 1920 |
| _ | | _ | _ | _ | | _ | | _ | | _ | ctc Leu | - | | | | 1968 |
| | | | | | _ | | | | | _ | gct Ala | _ | _ | | _ | 2016 |
| | | | | | | | | | | | atg Met | | | | | 2064 |

| | gac Asp 690 | | | | | | | | | | | | | | | 2112 |
|---|-------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|------|
| | cct Pro | _ | | _ | | | | | - | | | | | | | 2160 |
| _ | tct Ser | _ | | | - | | | | | | - | _ | | _ | | 2208 |
| | ttc Phe | | | | | | | | | | | | | | | 2256 |
| | gtt Val | | | | | | | | | | | | | | | 2304 |
| | tgc Cys 770 | | | | | | | | | | | | | | | 2352 |
| | cag Gln | | | | | | | | | | | | | | | 2400 |
| _ | gtg Val | | | _ | | _ | | | | | - | | _ | | _ | 2448 |
| | gaa Glu | | | | | | | | | | | | | | | 2496 |
| | gag Glu | _ | | _ | | _ | | | | | | | | | | 2544 |
| | cac His 850 | | | | | | | | | | | | | | | 2592 |
| | gcc Ala | | _ | | _ | | | | _ | | | _ | | | _ | 2640 |
| _ | tct Ser | _ | | | _ | | _ | _ | _ | _ | | _ | | | | 2688 |
| _ | gac Asp | | | | _ | | | | _ | | | _ | | _ | | 2736 |
| - | cca Pro | | | _ | | | | | | | - | | - | _ | - | 2784 |

| ctg ccc gtt tgg gcg cgg ccg gac tat aac ccc ccg cta gtg gag acg Leu Pro Val Trp Ala Arg Pro Asp Tyr Asn Pro Pro Leu Val Glu Thr 930 935 940 | 2832 |
|---|------|
| tgg aaa aag ccc gac tac gaa cca cct gtg gtc cat ggc aga aag acc Trp Lys Lys Pro Asp Tyr Glu Pro Pro Val Val His Gly Arg Lys Thr 945 950 955 960 | 2880 |
| aaa cgt aac acc aac cgg cgg ccg cag gac gtc aag ttc ccg ggt ggc Lys Arg Asn Thr Asn Arg Arg Pro Gln Asp Val Lys Phe Pro Gly Gly 965 970 975 | 2928 |
| ggt cag atc gtt ggt cgc agg ggc cct cct atc ccc aag gct cgt cgg Gly Gln Ile Val Gly Arg Arg Gly Pro Pro Ile Pro Lys Ala Arg Arg 980 985 990 | 2976 |
| ccc gag ggc agg acc tgg gct cag ccc ggt tac cct tgg ccc ctc tat Pro Glu Gly Arg Thr Trp Ala Gln Pro Gly Tyr Pro Trp Pro Leu Tyr 995 1000 1005 | 3024 |
| ggc aat aag gac aga cgg tct aca ggt aag tcc tgg ggt aag cca ggg Gly Asn Lys Asp Arg Arg Ser Thr Gly Lys Ser Trp Gly Lys Pro Gly 1010 1015 1020 | 3072 |
| tac cct tgg cca aga aag acc aaa cgt aac acc aac cga cgg ccg cag Tyr Pro Trp Pro Arg Lys Thr Lys Arg Asn Thr Asn Arg Arg Pro Gln 1025 1030 1035 1040 | 3120 |
| gac gtc aag ttc ccg ggt ggc ggt cag atc gtt ggt cgc agg ggc cct Asp Val Lys Phe Pro Gly Gly Gly Gln Ile Val Gly Arg Arg Gly Pro 1045 1050 1055 | 3168 |
| cct atc ccc aag gct cgt cgg ccc gag ggc agg acc tgg gct cag ccc Pro Ile Pro Lys Ala Arg Arg Pro Glu Gly Arg Thr Trp Ala Gln Pro 1060 1065 1070 | 3216 |
| ggt tac cct tgg ccc ctc tat ggc aat aag gac aga cgg tct acc ggt Gly Tyr Pro Trp Pro Leu Tyr Gly Asn Lys Asp Arg Arg Ser Thr Gly 1075 1080 1085 | 3264 |
| aag too tgg ggt aag coa ggg tat oot tgg coo Lys Ser Trp Gly Lys Pro Gly Tyr Pro Trp Pro 1090 1095 | 3297 |
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| <220> <223> MEFA 7.1 amino acid sequence | |
| <pre><400> 6 Met Ala Thr Lys Ala Val Cys Val Leu Lys Gly Asp Gly Pro Val Gln</pre> | |
| Gly Ile Ile Asn Phe Glu Gln Lys Glu Ser Asn Gly Pro Val Lys Val 20 25 30 | |

- Trp Gly Ser Ile Lys Gly Leu Thr Glu Gly Leu His Gly Phe His Val 35 40 45
- His Glu Phe Gly Asp Asn Thr Ala Gly Cys Thr Ser Ala Gly Pro His 50 55 60
- Phe Asn Pro Leu Ser Arg Lys His Gly Gly Pro Lys Asp Glu Glu Arg 65 70 75 80
- His Val Gly Asp Leu Gly Asn Val Thr Ala Asp Lys Asp Gly Val Ala $85 \\ 90 \\ 95$
- Asp Val Ser Ile Glu Asp Ser Val Ile Ser Leu Ser Gly Asp His Cys 100 105 110
- Ile Ile Gly Arg Thr Leu Val Val His Glu Lys Ala Asp Asp Leu Gly
 115 120 125
- Lys Gly Gly Asn Glu Glu Ser Thr Lys Thr Gly Asn Ala Gly Ser Arg 130 135 140
- Asn Cys Ser Ile Tyr Pro Gly His Ile Thr Gly His Arg Met Ala Trp 165 170 175
- Lys Leu Gly Ser Ala Ala Arg Thr Thr Ser Gly Phe Val Ser Leu Phe 180 185 190
- Ala Pro Gly Ala Lys Gln Asn Glu Thr His Val Thr Gly Gly Ala Ala 195 200 205
- Ala Arg Thr Thr Ser Gly Leu Thr Ser Leu Phe Ser Pro Gly Ala Ser 210 215 220
- Gln Asn Ile Gln Leu Ile Val Asp Phe Ile Pro Val Glu Asn Leu Glu 225 230 235 240
- Thr Thr Met Arg Ser Pro Val Phe Thr Asp Asn Ser Ser Pro Pro Val
 245 250 255
- Val Pro Gln Ser Phe Gln Val Ala His Leu His Ala Pro Thr Gly Ser 260 265 270
- Gly Lys Ser Thr Lys Val Pro Ala Ala Tyr Ala Ala Gln Gly Tyr Lys 275 280 285
- Val Leu Val Leu Asn Pro Ser Val Ala Ala Thr Leu Gly Phe Gly Ala 290 295 300

Tyr Met Ser Lys Ala His Gly Ile Asp Pro Asn Ile Arg Thr Gly Val Arg Thr Ile Thr Thr Gly Ser Pro Ile Thr Tyr Ser Thr Tyr Gly Lys 325 330 Phe Leu Ala Asp Gly Gly Cys Ser Gly Gly Ala Tyr Asp Ile Ile Ile Cys Asp Glu Cys His Ser Thr Asp Ala Thr Ser Ile Leu Gly Ile Gly Thr Val Leu Asp Gln Ala Glu Thr Ala Gly Ala Arg Leu Val Val Leu 375 Ala Thr Ala Thr Pro Pro Gly Ser Val Thr Val Pro His Pro Asn Ile 390 395 Glu Glu Val Ala Leu Ser Thr Thr Gly Glu Ile Pro Phe Tyr Gly Lys 405 Ala Ile Pro Leu Glu Val Ile Lys Gly Gly Arg His Leu Ile Phe Cys 425 His Ser Lys Lys Lys Cys Asp Glu Leu Ala Ala Lys Leu Val Ala Leu Gly Ile Asn Ala Val Ala Tyr Tyr Arg Gly Leu Asp Val Ser Val Ile Pro Thr Ser Gly Asp Val Val Val Ala Thr Asp Ala Leu Met Thr 470 475 Gly Tyr Thr Gly Asp Phe Asp Ser Val Ile Asp Cys Asn Thr Cys Val Thr Gln Thr Val Asp Phe Ser Leu Asp Pro Thr Phe Thr Ile Glu Thr 505 Ile Thr Leu Pro Gln Asp Ala Val Ser Arg Thr Gln Arg Arg Gly Arg Thr Gly Arg Gly Lys Pro Gly Ile Tyr Arg Phe Val Ala Pro Gly Glu 535 Arg Pro Ser Gly Met Phe Asp Ser Ser Val Leu Cys Glu Cys Tyr Asp 545 555 Ala Gly Cys Ala Trp Tyr Glu Leu Thr Pro Ala Glu Thr Thr Val Arq 570 Leu Arg Ala Tyr Met Asn Thr Pro Gly Leu Pro Val Cys Gln Asp His 580 Leu Glu Phe Trp Glu Gly Val Phe Thr Gly Leu Thr His Ile Asp Ala

- His Phe Leu Ser Gln Thr Lys Gln Ser Gly Glu Asn Leu Pro Tyr Leu 610 620
- Val Ala Tyr Gln Ala Thr Val Cys Ala Arg Ala Gln Ala Pro Pro 625 630 635 640
- Ser Trp Asp Gln Met Trp Lys Cys Leu Ile Arg Leu Lys Pro Thr Leu 645 650 655
- His Gly Pro Thr Pro Leu Leu Tyr Arg Leu Gly Ala Val Gln Asn Glu 660 665 670
- Ile Thr Leu Thr His Pro Val Thr Lys Tyr Ile Met Thr Cys Met Ser 675 680 685
- Ala Asp Leu Glu Val Val Thr Ser Ala Cys Ser Gly Lys Pro Ala Ile 690 695 700
- Ile Pro Asp Arg Glu Val Leu Tyr Arg Glu Phe Asp Glu Met Glu Glu 705 710 715 720
- Cys Ser Gln His Leu Pro Tyr Ile Glu Gln Gly Met Met Leu Ala Glu
 725 730 735
- Gln Phe Lys Gln Lys Ala Leu Gly Leu Ser Arg Gly Gly Lys Pro Ala
 740 745 750
- Ile Val Pro Asp Lys Glu Val Leu Tyr Gln Gln Tyr Asp Glu Met Glu 755 760 765
- Glu Cys Ser Gln Ala Ala Pro Tyr Ile Glu Gln Ala Gln Val Ile Ala 770 780
- His Gln Phe Lys Glu Lys Val Leu Gly Leu Ile Asp Asn Asp Gln Val
 785 790 795 800
- Val Val Thr Pro Asp Lys Glu Ile Leu Tyr Glu Ala Phe Asp Glu Met 805 810 815
- Glu Glu Cys Ala Ser Lys Ala Ala Leu Ile Glu Glu Gly Gln Arg Met 820 825 830
- Ala Glu Met Leu Lys Ser Lys Ile Gln Gly Leu Leu Gly Ile Leu Arg 835 840 845
- Arg His Val Gly Pro Gly Glu Gly Ala Val Gln Trp Met Asn Arg Leu 850 855 860
- Ile Ala Phe Ala Ser Arg Gly Asn His Val Ser Pro Thr His Tyr Val 865 870 875 880
- Pro Ser Arg Ser Arg Phe Ala Gln Ala Leu Pro Val Trp Ala Arg 885 890 895
- Pro Asp Tyr Asn Pro Pro Leu Val Glu Thr Trp Lys Lys Pro Asp Tyr 900 905 910

Glu Pro Pro Val Val His Gly Arg Ser Ser Arg Arg Phe Ala Gln Ala 915 920 925

Leu Pro Val Trp Ala Arg Pro Asp Tyr Asn Pro Pro Leu Val Glu Thr 930 935 940

Trp Lys Lys Pro Asp Tyr Glu Pro Pro Val Val His Gly Arg Lys Thr 945 950 955 960

Lys Arg Asn Thr Asn Arg Arg Pro Gln Asp Val Lys Phe Pro Gly Gly 965 970 975

Gly Gln Ile Val Gly Arg Arg Gly Pro Pro Ile Pro Lys Ala Arg Arg 980 985 990

Pro Glu Gly Arg Thr Trp Ala Gln Pro Gly Tyr Pro Trp Pro Leu Tyr 995 1000 1005

Gly Asn Lys Asp Arg Arg Ser Thr Gly Lys Ser Trp Gly Lys Pro Gly 1010 1015 1020

Tyr Pro Trp Pro Arg Lys Thr Lys Arg Asn Thr Asn Arg Arg Pro Gln 1025 1030 1035 1040

Asp Val Lys Phe Pro Gly Gly Gly Gln Ile Val Gly Arg Arg Gly Pro 1045 1050 1055

Pro Ile Pro Lys Ala Arg Arg Pro Glu Gly Arg Thr Trp Ala Gln Pro 1060 1065 1070

Gly Tyr Pro Trp Pro Leu Tyr Gly Asn Lys Asp Arg Arg Ser Thr Gly
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Lys Ser Trp Gly Lys Pro Gly Tyr Pro Trp Pro 1090 1095

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Gly Ala Lys Gln Asn

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